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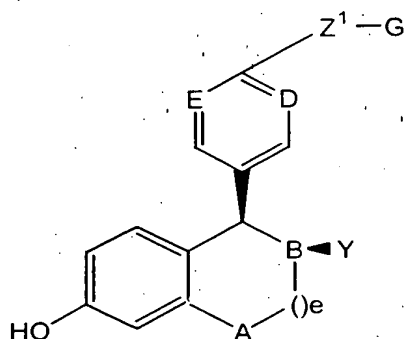
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Claims

1. A process for preparing a compound of the formula:



wherein:

A is selected from CH_2 and NR ;

B, D and E are independently selected from CH and N;

Y is

- (a) phenyl, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (b) naphthyl, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (c) $\text{C}_3\text{-C}_8$ cycloalkyl, optionally substituted with 1-2 substituents independently selected from R^4 ;
- (d) $\text{C}_3\text{-C}_8$ cycloalkynyl, optionally substituted with 1-2 substituents independently selected from R^4 ;
- (e) a five membered heterocycle containing up to two heteroatoms selected from the group consisting of $-\text{O}-$, $-\text{NR}^2-$ and $-\text{S}(\text{O})_n-$, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (f) a six membered heterocycle containing up to two heteroatoms selected from the group consisting of $-\text{O}-$, $-\text{NR}^2-$ and $-\text{S}(\text{O})_n-$ optionally substituted with 1-3 substituents independently selected from R^4 ; or
- (g) a bicyclic ring system consisting of a five or six membered heterocyclic ring fused to a phenyl ring, said heterocyclic ring containing up to two heteroatoms selected from the group consisting of $-\text{O}-$, $-\text{NR}^2-$, NR^2- and $-\text{S}(\text{O})_n-$, optionally

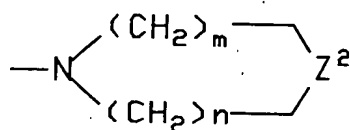
substituted with 1-3 substituents independently selected from R^4 ;

Z^1 is

- (a) $-(CH_2)_p W(CH_2)_q-$;
- (b) $-O(CH_2)_p CR^5R^6-$;
- (c) $-O(CH_2)_p W(CH_2)_q$;
- (d) $-OCHR^2CHR^3-$; or
- (e) $-SCHR^2CHR^3-$;

G is

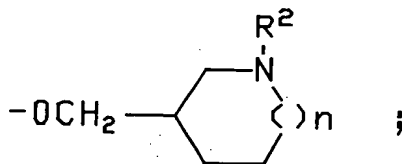
- (a) $-NR^7R^8$;
- (b)



wherein n is 0, 1 or 2; m is 1, 2 or 3; Z^2 is $-NH-$, $-O-$, $-S-$, or $-CH_2-$; optionally fused on adjacent carbon atoms with one or two phenyl rings and, optionally independently substituted on carbon with one to three substituents and, optionally, independently on nitrogen with a chemically suitable substituent selected from R^4 ; or

- (c) a bicyclic amine containing five to twelve carbon atoms, either bridged or fused and optionally substituted with 1-3 substituents independently selected from R^4 ;

Z^1 and G in combination may be



W is

- (a) $-CH_2-$;
- (b) $-CH=CH-$;
- (c) $-O-$;
- (d) $-NR^2-$;

(e) $-S(O)_n-$;

(f)

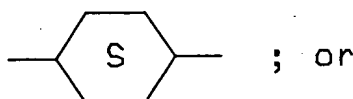


(g) $-CR^2(OH)-$;

(h) $-CONR^2-$;

(i) $-NR^2CO-$;

(j)



(k) $-C\equiv C-$;

R is hydrogen or C_1-C_6 alkyl;

R^2 and R^3 are independently

(a) hydrogen; or

(b) C_1-C_4 alkyl;

R^4 is

(a) hydrogen;

(b) halogen;

(c) C_1-C_6 alkyl;

(d) C_1-C_4 alkoxy;

(e) C_1-C_4 acyloxy;

(f) C_1-C_4 alkylthio;

(g) C_1-C_4 alkylsulfinyl;

(h) C_1-C_4 alkylsulfonyl;

(i) hydroxy (C_1-C_4)alkyl;

(j) aryl (C_1-C_4)alkyl;

(k) $-CO_2H$;

(l) $-CN$;

(m) $-CONHOR$;

(n) $-SO_2NHR$;

(o) $-NH_2$;

(p) C_1-C_4 alkylamino;

- (q) C₁-C₄ dialkylamino;
- (r) -NHSO₂R;
- (s) -NO₂;
- (t) -aryl; or
- (u) -OH.

R⁵ and R⁶ are independently C₁-C₈ alkyl or together form a C₃-C₁₀ carbocyclic ring;

R⁷ and R⁸ are independently

- (a) phenyl;
- (b) a C₃-C₁₀ carbocyclic ring, saturated or unsaturated;
- (c) a C₃-C₁₀ heterocyclic ring containing up to two heteroatoms, selected from -O-, -N- and -S-;
- (d) H;
- (e) C₁-C₆ alkyl; or
- (f) form a 3 to 8 membered nitrogen containing ring with R⁵ or R⁶;

R⁷ and R⁸ in either linear or ring form may optionally be substituted with up to three substituents independently selected from C₁-C₆ alkyl, halogen, alkoxy, hydroxy and carboxy;

a ring formed by R⁷ and R⁸ may be optionally fused to a phenyl ring;

e is 0, 1 or 2;

m is 1, 2 or 3;

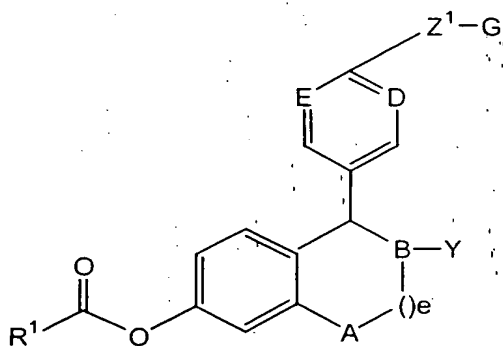
n is 0, 1 or 2;

p is 0, 1, 2 or 3;

q is 0, 1, 2 or 3;

and optical and geometric isomers thereof;

comprising selectively deacetylating a compound of the formula



wherein R¹ is (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl wherein the alkyl, alkenyl or alkynyl groups are optionally substituted by one to three halo, in the presence of a hydrolytic enzyme and an aqueous buffer solution.

- 5 2. A process according to claim 1, wherein the hydrolytic enzyme is
lipase.
3. A process according to claim 1, wherein the hydrolytic enzyme is
esterase.
4. A process according to claim 1, wherein the hydrolytic enzyme is liver
10 acetone powder.
5. A process according to claim 1, wherein the hydrolytic enzyme is
lipase from *Porcine pancreas*, cholesterol esterase from *Pseudomonas Fluorescens*
and cholesterol esterase from *Porcine pancreas*.
6. A process according to claim 2, wherein the lipase is GC-4, PS30,
15 AY30, PGE, AK, N, L-10, AP-12, FAP-15, R-10, G, MAP10, SAM II, lipase from
Pseudomonas fluorescens, lipase from *Candida cylindracea*, Lip-300, lipase from
Chromobacterium viscosum, lipase from *Mucor miehei*, lipase from *Pancreatic*, lipase
from *Pseudomonas fluorescens*, lipase from *Rhizopus niveus*, PPL, type II, lipase
from *Wheat germ*, lipase from *Rhizopus arrhizus*, lipase from *Mucor javanicus*, lipase
20 from *Pseudomonas cepacia*, lipase from *Cadia lipolytica*, lipase from *Penicillium*
roqueforti, lipoprotein lipase ca#70-6571-01, lipase from *Porcine pancreas*, and
lipoprotein lipase ca# 70-1481-01.
7. A process according to claim 3, wherein the esterase is PLE-A,
immobilized, hog liver, esterase from *Hog pancreas*, *Porcine liver* E-3128,
25 cholesterin-esterase, cholesterol esterase from *Pseudomonas fluorescens*,
cholesterol esterase from *Porcine pancreas*, cholesterol esterase from *Bovine*
pancreas, cholesterol esterase from *Pseudomonas fluorescens*, cholesterol esterase

from *Porcine liver*, cholesterol esterase from *Rabbit liver*, cholinesterase, cholinesterase from *Electric eel*, cholinesterase, choloylglycine hydrolase, esterase from *Thermoanaerobium Brockii*, esterase from *Bacillus sp* and esterase from *Mucor miehi*.

5 8. A process according to claim 4, wherein the liver acetone powder is cat I-1256, dog I7379, eel I-1266, horse I9627, calf I7876, guinea pig I1631, mouse I8254, goat I2635, chicken I8001, sheep I0132, pigeon I8376, seal I7627, rattlesnake I9885, trout I5131, turtle I-0757, rat I1380, lungfish I7377, salmon I7502, eel (electrophorus electricus) I8380 and lemon shark I1130.

10 9. A process according to claim 1, wherein the hydrolytic enzyme is immobilized on a solid support.

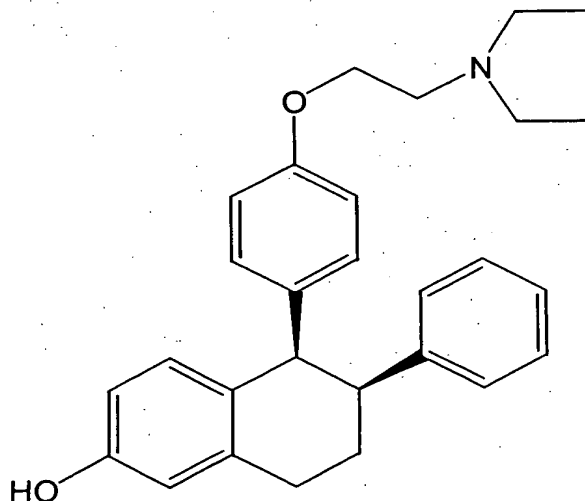
 10. A process according to claim 1, wherein the hydrolytic enzyme is a cross-linked enzyme.

15 11. A process according to claim 1, wherein the lipase is in pure crystalline form.

 12. A process according to claim 1, wherein the aqueous buffer solution is a phosphate, citric acid or boronic acid solution.

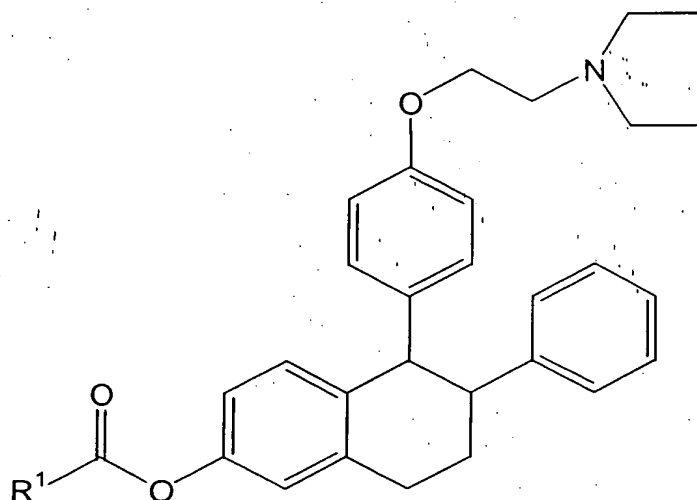
 13. A process according to claim 1, wherein the aqueous buffer solution has a pH between a pH of about 6 to a pH of about 8.

20 14. A process according to claim 1, for preparing a compound of the formula



VII

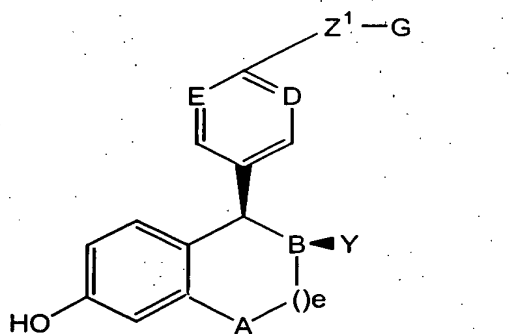
comprising selectively deacetylating a compound of the formula



VIII

wherein R^1 is (C_1-C_6) alkyl, (C_2-C_6) alkynyl, (C_2-C_6) alkenyl wherein the alkyl, alkenyl or alkynyl groups are optionally substituted by one to three halo in the presence of a hydrolytic enzyme and an aqueous buffer solution.

15. A process for preparing a compound of the formula:



10

15 wherein:

A is selected from CH_2 and NR ;

B, D and E are independently selected from CH and N;

Y is

- (a) phenyl, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (b) naphthyl, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (c) C_3-C_8 cycloalkyl, optionally substituted with 1-2 substituents independently selected from R^4 ;

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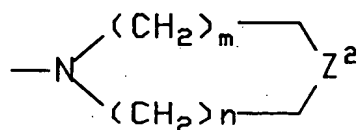
- (d) ¹⁰ C₃-C₈ cycloalkynyl, optionally substituted with 1-2 substituents independently selected from R⁴;
- (e) a five membered heterocycle containing up to two heteroatoms selected from the group consisting of -O-, -NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴;
- (f) a six membered heterocycle containing up to two heteroatoms selected from the group consisting of -O-, -NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴; or
- (g) a bicyclic ring system consisting of a five or six membered heterocyclic ring fused to a phenyl ring, said heterocyclic ring containing up to two heteroatoms selected from the group consisting of -O-, -NR²-, NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴;

Z¹ is

- (a) -(CH₂)_p W(CH₂)_q-;
- (b) -O(CH₂)_p CR⁵R⁶-;
- (c) -O(CH₂)_p W(CH₂)_q;
- (d) -OCHR²CHR³-; or
- (e) -SCHR²CHR³-;

G is

- (a) -NR⁷R⁸;
- (b)

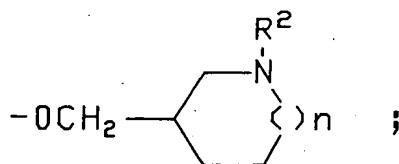


wherein n is 0, 1 or 2; m is 1, 2 or 3; Z² is -NH-, -O-, -S-, or -CH₂-; optionally fused on adjacent carbon atoms with one or two phenyl rings and, optionally independently substituted on carbon with one to three substituents and, optionally,

independently on nitrogen with a chemically suitable substituent selected from R^4 ; or

- (c) a bicyclic amine containing five to twelve carbon atoms, either bridged or fused and optionally substituted with 1-3 substituents independently selected from R^4 ;

Z^1 and G in combination may be

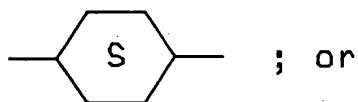


W is

- (a) $-CH_2-$;
 (b) $-CH=CH-$;
 (c) $-O-$;
 (d) $-NR^2-$;
 (e) $-S(O)_n-$;
 (f)



- (g) $-CR^2(OH)-$;
 (h) $-CONR^2-$;
 (i) $-NR^2CO-$;
 (j)



- (k) $-C\equiv C-$;

R is hydrogen or C_1 - C_6 alkyl;

R^2 and R^3 are independently

- (a) hydrogen; or
 (b) C_1 - C_4 alkyl;

R^4 is

- (a) hydrogen;

- (b) halogen;
- (c) C₁-C₆ alkyl;
- (d) C₁-C₄ alkoxy;
- (e) C₁-C₄ acyloxy;
- (f) C₁-C₄ alkylthio;
- (g) C₁-C₄ alkylsulfinyl;
- (h) C₁-C₄ alkylsulfonyl;
- (i) hydroxy (C₁-C₄)alkyl;
- (j) aryl (C₁-C₄)alkyl;
- (k) -CO₂H;
- (l) -CN;
- (m) -CONHOR;
- (n) -SO₂NHR;
- (o) -NH₂;
- (p) C₁-C₄ alkylamino;
- (q) C₁-C₄ dialkylamino;
- (r) -NHSO₂R;
- (s) -NO₂;
- (t) -aryl; or
- (u) -OH.

R⁵ and R⁶ are independently C₁-C₈ alkyl or together form a C₃-C₁₀ carbocyclic ring;

R⁷ and R⁸ are independently

- (a) phenyl;
- (b) a C₃-C₁₀ carbocyclic ring, saturated or unsaturated;
- (c) a C₃-C₁₀ heterocyclic ring containing up to two heteroatoms, selected from -O-, -N- and -S-;
- (d) H;
- (e) C₁-C₆ alkyl; or
- (f) form a 3 to 8 membered nitrogen containing ring with R⁵ or R⁶;

R⁷ and R⁸ in either linear or ring form may optionally be substituted with up to three substituents independently selected from C₁-C₆ alkyl, halogen, alkoxy, hydroxy and carboxy;

a ring formed by R⁷ and R⁸ may be optionally fused to a phenyl ring;

e is 0, 1 or 2;

m is 1, 2 or 3;

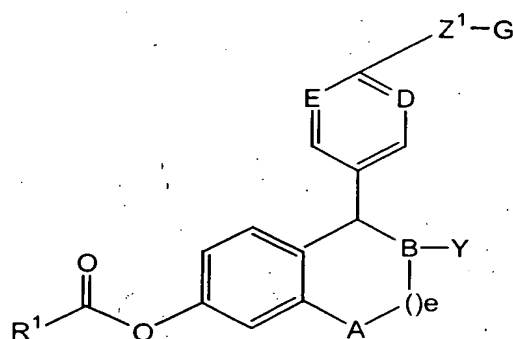
n is 0, 1 or 2;

p is 0, 1, 2 or 3;

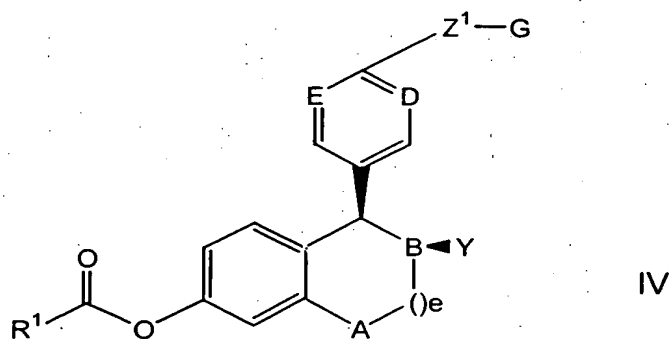
q is 0, 1, 2 or 3;

and optical and geometric isomers thereof;

comprising enzymatically resolving of a compound of the formula



- 10 wherein R¹ is (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl wherein the alkyl, alkenyl or alkynyl groups are optionally substituted by one to three halo in the presence of a lipase and an aqueous buffer solution; and (b) reacting the compound of formula IV so formed



- 15 wherein R¹ is as defined above, with a base in the presence of a polar protic solvent.

16. A process according to claim 15, wherein the aqueous buffer solution is a phosphate, citric acid or boronic acid solution.

17. A process according to claim 15, wherein the lipase from *Mucor miehei*.

18. A process according to claim 15, wherein the base is sodium methoxy, sodium hydroxide, lithium hydroxide or potassium hydroxide.

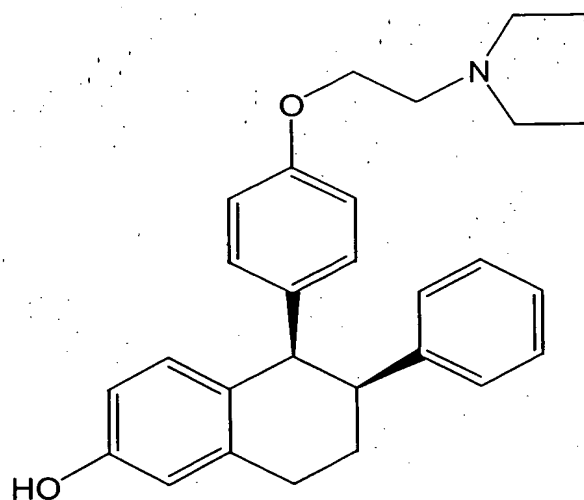
19. A process according to claim 15, wherein the polar protic solvent is methanol, ethanol or water.

20. A process according to claim 15, wherein the lipase is immobilized on a solid support.

21. A process according to claim 15, wherein the lipase is a cross-linked enzyme.

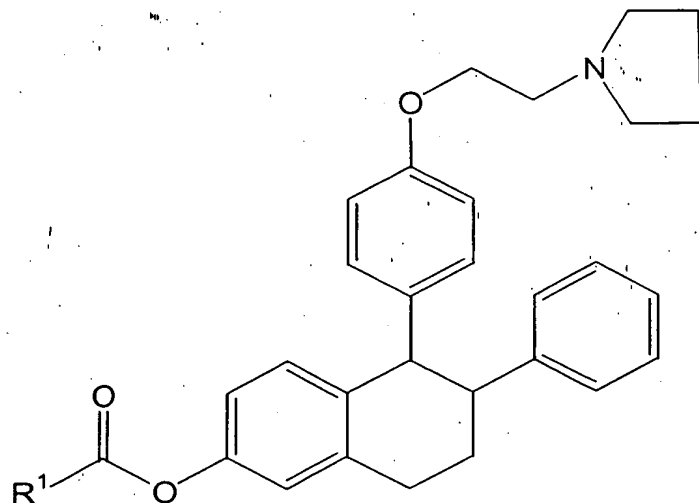
22. A process according to claim 15, wherein the lipase is in pure crystalline form.

23. A process according to claim 15, for preparing a compound of the formula



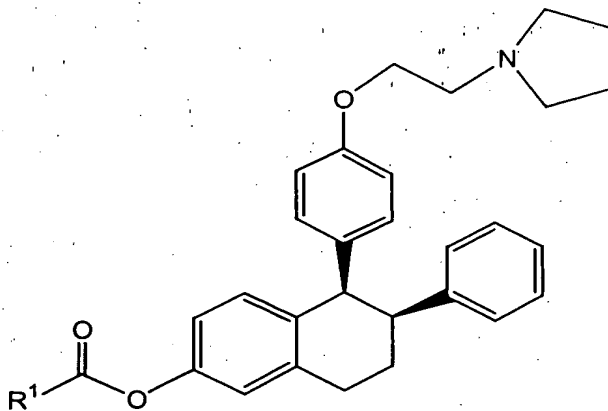
VII

comprising enzymatically resolving of a compound of the formula



VIII

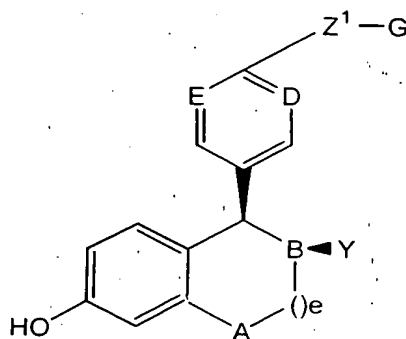
- 5 wherein R¹ is (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl wherein the alkyl, alkenyl or alkynyl groups are optionally substituted by one to three halo in the presence of a lipase and an aqueous buffer solution; and (b) reacting the compound of Formula X so formed



X

wherein R¹ is as defined above, with a base in the presence of a polar protic solvent.

24. A process for preparing a compound of the formula:



wherein:

A is selected from CH_2 and NR ;

B, D and E are independently selected from CH and N;

Y is

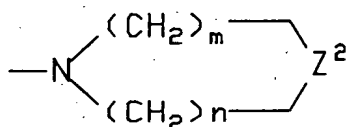
- (a) phenyl, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (b) naphthyl, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (c) C_3 - C_8 cycloalkyl, optionally substituted with 1-2 substituents independently selected from R^4 ;
- (d) C_3 - C_8 cycloalkynyl, optionally substituted with 1-2' substituents independently selected from R^4 ;
- (e) a five membered heterocycle containing up to two heteroatoms selected from the group consisting of $-\text{O}-$, $-\text{NR}^2$ - and $-\text{S}(\text{O})_n$ -, optionally substituted with 1-3 substituents independently selected from R^4 ;
- (f) a six membered heterocycle containing up to two heteroatoms selected from the group consisting of $-\text{O}-$, $-\text{NR}^2$ - and $-\text{S}(\text{O})_n$ -, optionally substituted with 1-3 substituents independently selected from R^4 ; or
- (g) a bicyclic ring system consisting of a five or six membered heterocyclic ring fused to a phenyl ring, said heterocyclic ring containing up to two heteroatoms selected from the group consisting of $-\text{O}-$, $-\text{NR}^2$ -, NR^2 - and $-\text{S}(\text{O})_n$ -, optionally substituted with 1-3 substituents independently selected from R^4 ;

Z¹ is

- (a) $-(CH_2)_p W(CH_2)_q-$;
- (b) $-O(CH_2)_p CR^5R^6-$;
- (c) $-O(CH_2)_p W(CH_2)_q$;
- (d) $-OCHR^2CHR^3-$; or
- (e) $-SCHR^2CHR^3-$;

G is

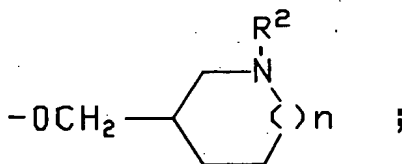
- (a) $-NR^7R^8$;
- (b)



wherein n is 0, 1 or 2; m is 1, 2 or 3; Z² is -NH-, -O-, -S-, or -CH₂-; optionally fused on adjacent carbon atoms with one or two phenyl rings and, optionally independently substituted on carbon with one to three substituents and, optionally, independently on nitrogen with a chemically suitable substituent selected from R⁴; or

- (c) a bicyclic amine containing five to twelve carbon atoms, either bridged or fused and optionally substituted with 1-3 substituents independently selected from R⁴;

Z¹ and G in combination may be

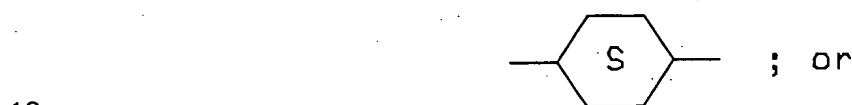


W is

- (a) $-CH_2-$;
- (b) $-CH=CH-$;
- (c) $-O-$;
- (d) $-NR^2-$;
- (e) $-S(O)_n-$;
- (f)



- 5 (g) $-\text{CR}^2(\text{OH})-$;
 (h) $-\text{CONR}^2-$;
 (i) $-\text{NR}^2\text{CO}-$;
 (j)



- (k) $-\text{C}\equiv\text{C}-$;

R is hydrogen or $\text{C}_1\text{-C}_6$ alkyl;

R^2 and R^3 are independently

- 15 (a) hydrogen; or
 (b) $\text{C}_1\text{-C}_4$ alkyl;

R^4 is

- 20 (a) hydrogen;
 (b) halogen;
 (c) $\text{C}_1\text{-C}_6$ alkyl;
 (d) $\text{C}_1\text{-C}_4$ alkoxy;
 (e) $\text{C}_1\text{-C}_4$ acyloxy;
 (f) $\text{C}_1\text{-C}_4$ alkylthio;
 (g) $\text{C}_1\text{-C}_4$ alkylsulfinyl;
 (h) $\text{C}_1\text{-C}_4$ alkylsulfonyl;
 25 (i) hydroxy ($\text{C}_1\text{-C}_4$)alkyl;
 (j) aryl ($\text{C}_1\text{-C}_4$)alkyl;
 (k) $-\text{CO}_2\text{H}$;
 (l) $-\text{CN}$;
 (m) $-\text{CONHOR}$;
 30 (n) $-\text{SO}_2\text{NHR}$;
 (o) $-\text{NH}_2$;
 (p) $\text{C}_1\text{-C}_4$ alkylamino;
 (q) $\text{C}_1\text{-C}_4$ dialkylamino;
 (r) $-\text{NHSO}_2\text{R}$;

- (s) -NO₂;
- (t) -aryl; or
- (u) -OH.

R⁵ and R⁶ are independently C₁-C₈ alkyl or together form a C₃-C₁₀ carbocyclic ring;

R⁷ and R⁸ are independently

- (a) phenyl;
- (b) a C₃-C₁₀ carbocyclic ring, saturated or unsaturated;
- (c) a C₃-C₁₀ heterocyclic ring containing up to two heteroatoms, selected from -O-, -N- and -S-;
- (d) H;
- (e) C₁-C₆ alkyl; or
- (f) form a 3 to 8 membered nitrogen containing ring with R⁵ or R⁶;

R⁷ and R⁸ in either linear or ring form may optionally be substituted with up to three substituents independently selected from C₁-C₆ alkyl, halogen, alkoxy, hydroxy and carboxy;

a ring formed by R⁷ and R⁸ may be optionally fused to a phenyl ring;

e is 0, 1 or 2;

m is 1, 2 or 3;

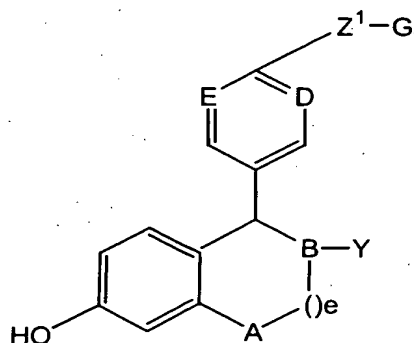
n is 0, 1 or 2;

p is 0, 1, 2 or 3;

q is 0, 1, 2 or 3;

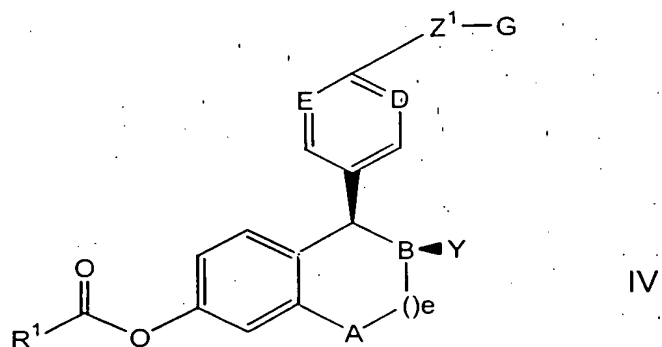
and optical and geometric isomers thereof;

comprising enzymatically resolving of a compound of the formula



VI

in the presence of a lipase and an acetylating agent, and (b) reacting the compound of formula IV so formed



wherein R¹ is (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl wherein the alkyl, alkenyl or alkynyl groups are optionally substituted by one to three halo, with a base in the presence of a polar protic solvent.

25. A process according to claim 24, wherein the hydrolytic enzyme is a lipase.

26. A process according to claim 24, wherein the lipase is GC-4, PS30, AY30, PGE, AK, N, L-10, AP-12, FAP-15, R-10, G, MAP10, SAM II, lipase from *Pseudomonas fluorescens*, lipase from *Candida cylindracea*, Lip-300, lipase from *Chromobacterium viscosum*, lipase from *Mucor miehei*, lipase from *Pancreatic*, lipase from *Pseudomonas fluorescens*, lipase from *Rhizopus niveus*, PPL, type II, lipase from *Wheat germ*, lipase from *Rhizopus arrhizus*, lipase from *Mucor javanicus*, lipase from *Pseudomonas cepacia*, lipase from *Cadia lipolytica*, lipase from *Penicillium roqueforti*, lipoprotein lipase ca#70-6571-01, lipase from *Porcine pancreas*, and lipoprotein lipase ca# 70-1481-01.

27. A process according to claim 24, wherein the acetylating agent is ethyl acetate, vinyl acetate, chloroacetate or trifluoroacetate.

28. A process according to claim 24, wherein the base is sodium methoxy, sodium hydroxide, lithium hydroxide or potassium hydroxide.

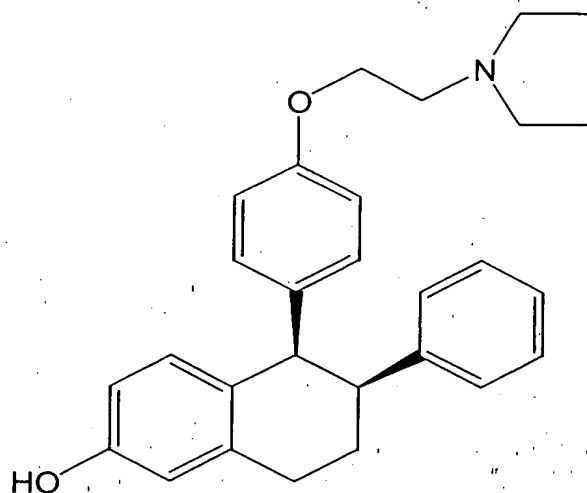
29. A process according to claim 24, wherein the polar protic solvent is methanol, ethanol or water.

30. A process according to claim 24, wherein the lipase is immobilized on a solid support.

31. A process according to claim 24, wherein the lipase is a cross-linked enzyme.

32. A process according to claim 24, wherein the lipase is in pure crystalline form.

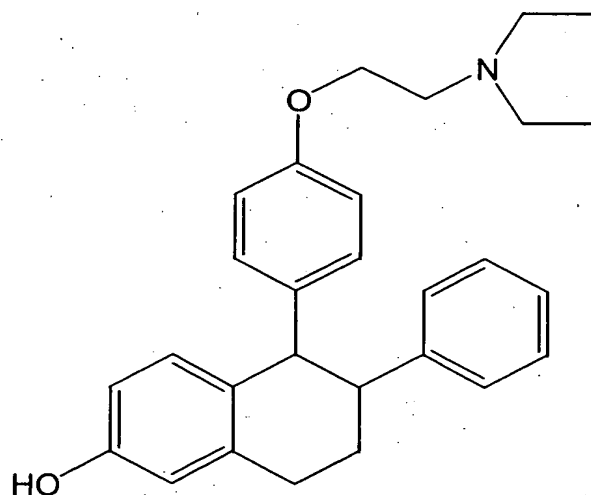
5 33. A process according to claim 24, for preparing a compound of the formula



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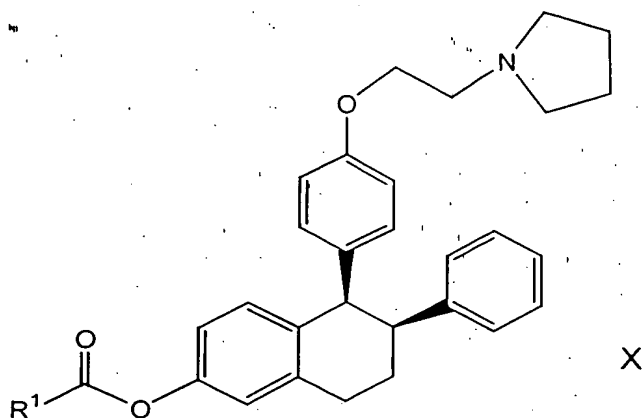
comprising enzymatically resolving of a compound of the formula

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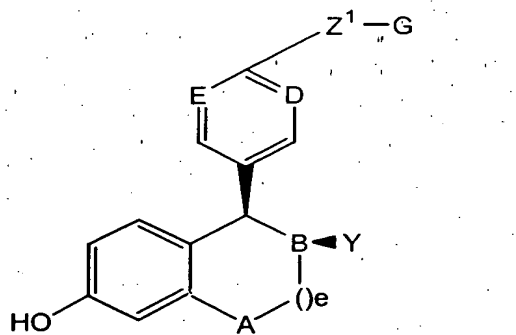
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in the presence of a lipase and acetylating agent, and (b) reacting the compound of Formula X so formed



wherein R¹ is (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl wherein the alkyl, alkenyl or alkynyl groups are optionally substituted by one to three halo, with a base in the presence of a polar protic solvent.

34. A process for preparing a compound of the formula:



wherein:

A is selected from CH₂ and NR;

B, D and E are independently selected from CH and N;

Y is

- (a) phenyl, optionally substituted with 1-3 substituents independently selected from R⁴;
- (b) naphthyl, optionally substituted with 1-3 substituents independently selected from R⁴;
- (c) C₃-C₈ cycloalkyl, optionally substituted with 1-2 substituents independently selected from R⁴;
- (d) C₃-C₈ cycloalkynyl, optionally substituted with 1-2 substituents independently selected from R⁴;

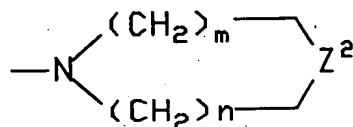
- (e) a five membered heterocycle containing up to two heteroatoms selected from the group consisting of -O-, -NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴;
- (f) a six membered heterocycle containing up to two heteroatoms selected from the group consisting of -O-, -NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴; or
- (g) a bicyclic ring system consisting of a five or six membered heterocyclic ring fused to a phenyl ring, said heterocyclic ring containing up to two heteroatoms selected from the group consisting of -O-, -NR²-, NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴;

Z¹ is

- (a) -(CH₂)_p W(CH₂)_q-;
- (b) -O(CH₂)_p CR⁵R⁶-;
- (c) -O(CH₂)_pW(CH₂)_q;
- (d) -OCHR²CHR³-; or
- (e) -SCHR²CHR³-;

G is

- (a) -NR⁷R⁸;
- (b)

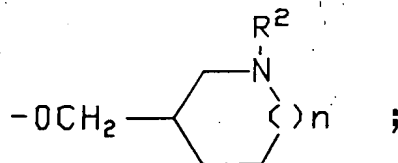


wherein n is 0, 1 or 2; m is 1, 2 or 3; Z² is -NH-, -O-, -S-, or -CH₂-, optionally fused on adjacent carbon atoms with one or two phenyl rings and, optionally independently substituted on carbon with one to three substituents and, optionally, independently on nitrogen with a chemically suitable substituent selected from R⁴; or

- (c) a bicyclic amine containing five to twelve carbon atoms, either bridged or fused and optionally substituted with 1-3 substituents independently selected from R^4 ;

Z_1 and G in combination may be

5



W is

10

- (a) $-\text{CH}_2-$;
- (b) $-\text{CH}=\text{CH}-$;
- (c) $-\text{O}-$;
- (d) $-\text{NR}^2-$;
- (e) $-\text{S}(\text{O})_n-$;
- (f)

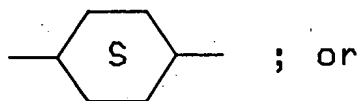
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- (g) $-\text{CR}^2(\text{OH})-$;
- (h) $-\text{CONR}^2-$;
- (i) $-\text{NR}^2\text{CO}-$;
- (j)

25



- (k) $-\text{C}\equiv\text{C}-$;

R is hydrogen or $\text{C}_1\text{-C}_6$ alkyl;

R^2 and R^3 are independently

30

- (a) hydrogen; or
- (b) $\text{C}_1\text{-C}_4$ alkyl;

R^4 is

- (a) hydrogen;
- (b) halogen;
- (c) $\text{C}_1\text{-C}_6$ alkyl;

- 5
- (d) C₁-C₄ alkoxy;
(e) C₁-C₄ acyloxy;
(f) C₁-C₄ alkylthio;
(g) C₁-C₄ alkylsulfinyl;
(h) C₁-C₄ alkylsulfonyl;
(i) hydroxy (C₁-C₄)alkyl;
(j) aryl (C₁-C₄)alkyl;
(k) -CO₂H;
(l) -CN;
10 (m) -CONHOR;
(n) -SO₂NHR;
(o) -NH₂;
(p) C₁-C₄ alkylamino;
(q) C₁-C₄ dialkylamino;
15 (r) -NHSO₂R;
(s) -NO₂;
(t) -aryl; or
(u) -OH.

20 R⁵ and R⁶ are independently C₁-C₈ alkyl or together form a C₃-C₁₀ carbocyclic ring;

R⁷ and R⁸ are independently

- 25 (a) phenyl;
(b) a C₃-C₁₀ carbocyclic ring, saturated or unsaturated;
(c) a C₃-C₁₀ heterocyclic ring containing up to two heteroatoms,
selected from -O-, -N- and -S-;
(d) H;
(e) C₁-C₆ alkyl; or
(f) form a 3 to 8 membered nitrogen containing ring with R⁵ or R⁶;

30 R⁷ and R⁸ in either linear or ring form may optionally be substituted with up to three substituents independently selected from C₁-C₆ alkyl, halogen, alkoxy, hydroxy and carboxy;

a ring formed by R⁷ and R⁸ may be optionally fused to a phenyl ring;

e is 0, 1 or 2;

m is 1, 2 or 3;

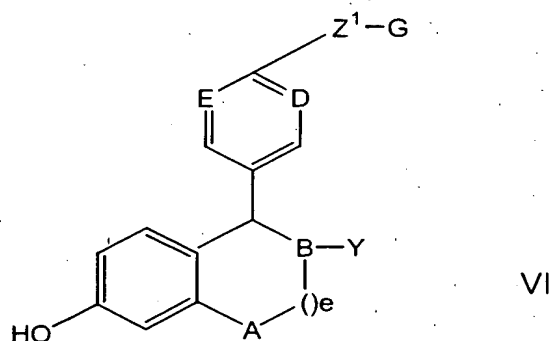
n is 0, 1 or 2;

p is 0, 1, 2 or 3;

q is 0, 1, 2 or 3;

and optical and geometric isomers thereof;

5 comprising enzymatically resolving of a compound of the formula



wherein R¹ is (C₁-C₆)alkyl, in the presence of lipase,

35. A process according to claim 34, wherein the lipase is *Mucor miehei*.

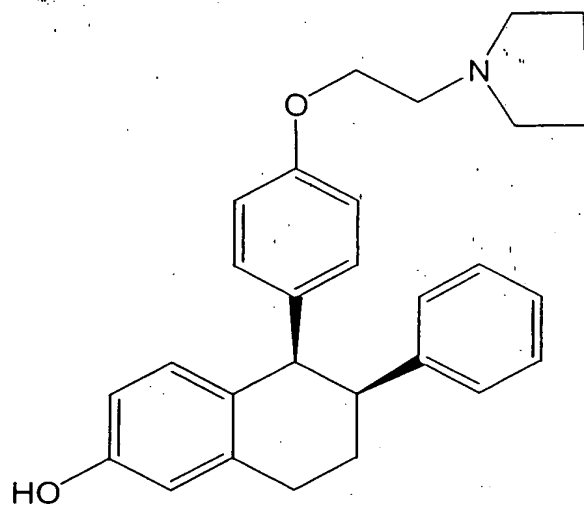
10 36. A process according to claim 34, wherein the lipase is immobilized on a solid support.

37. A process according to claim 34, wherein the lipase is a cross-linked enzyme.

15 38. A process according to claim 34, wherein the lipase is in pure crystalline form.

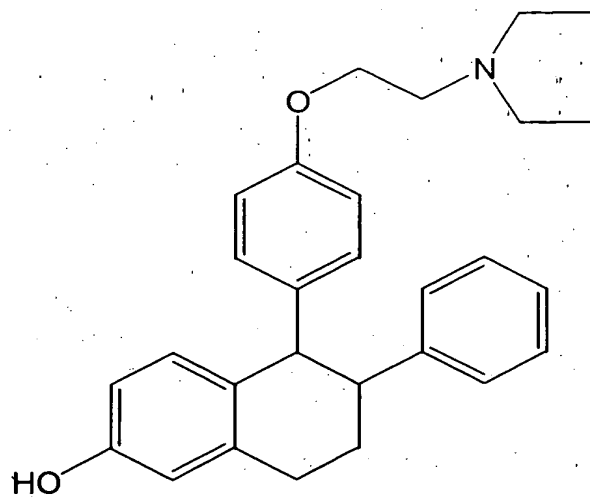
39. A process according to claim 34, wherein the hydrolytic enzyme is lipase from *Porcine pancreas*, cholesterol esterase from *Pseudomonas Fluoriscens* and cholesterol esterase from *Porcine pancreas*.

20 40. A process according to claim 34, for preparing a compound of the formula



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comprising enzymatically resolving of a compound of the formula



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in the presence of lipase.